

Attorney Docket No.: **PENN-0742**
Inventors: **Zhe Lu**
Serial No.: **09/743,054**
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Listing of Claims:

Claim 1 (original): A modified tertiapin peptide comprising
SEQ ID NO:2.

Claim 2 (original): A method of inhibiting activity of
inward-rectifier potassium channels comprising administering to
an animal a compound comprising a tertiapin-like α helix.

Claim 3 (original): The method of claim 2 wherein the
compound comprises tertiapin.

Claim 4 (original): The method of claim 2 wherein the
compound comprises SEQ ID NO:2.

Claim 5 (original): A method of identifying compounds
capable of inhibiting activity of inward-rectifier potassium
channels comprising:

(a) administering a test compound to an animal or cell
culture system;

(b) measuring activity of inward-rectifier potassium
channels in the animal or cell culture system;

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(c) and comparing the measured activity with a level of activity of the channels following administration of tertiapin or a modified tertiapin peptide of claim 1 to the animal or cell culture system, wherein a measured activity equal to or less than the levels of activity following administration of tertiapin or a modified tertiapin peptide of claim 1 is indicative of the test compound being an inhibitor.

Claim 6 (original): A method of identifying compounds capable of inhibiting activity of inward-rectifier potassium channels comprising:

(a) administering detectably labeled tertiapin or modified tertiapin peptide of claim 1 to a cell culture system, purified inward rectifier potassium channels or an animal;

(b) administering a test compound to the cell culture, purified inward-rectifier potassium channels or animal;

(c) and detecting unbound labeled tertiapin or the modified tertiapin peptide of claim 1 wherein the presence of unbound labeled tertiapin or the modified tertiapin peptide of claim 1 is indicative of the test compound being an inhibitor.

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Claim 7 (original): A pharmaceutical composition comprising a compound having a tertiapin-like α helix and a pharmaceutically acceptable vehicle.

Claim 8 (original): The pharmaceutical composition of claim 7 wherein the compound comprises SEQ ID NO:2.

Claim 9 (original): A method of controlling insulin secretion in a mammal comprising administering to the mammal a pharmaceutical composition of claim 7.

Claim 10 (original): A method of controlling cardiac rhythm and electrical conduction in a mammal comprising administering to the mammal a pharmaceutical composition of claim 7.

Claim 11 (original): A method of inducing diuresis in a mammal comprising administering to the mammal a pharmaceutical composition of claim 7.

Claim 12 (original): A method of modulating neurotransmissions in neurons of the nervous system of a mammal

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comprising administering to the mammal a pharmaceutical composition of claim 7.

Claim 13 (original): A method for rational design of drugs targeted to inward-rectifier K⁺ channels comprising:

(a) assessing distances of residues of tertiapin or SEQ ID NO:2 critical to binding of tertiapin or SEQ ID NO:2 to an inward-rectifier K⁺ channel;

(b) synthesizing a compound with residues at distance similar to those assess for tertiapin; and

(c) determining the ability of the compound to bind to inward rectifier K⁺ channels.

Claim 14 (original): A method for rational design of drugs targeted to inward-rectifier K⁺ channels comprising:

(a) synthesizing a compound having a similar structure or amino acid sequence or amino acid composition to tertiapin; and

(b) determining the ability of the compound to bind to inward-rectifier K⁺ channels.